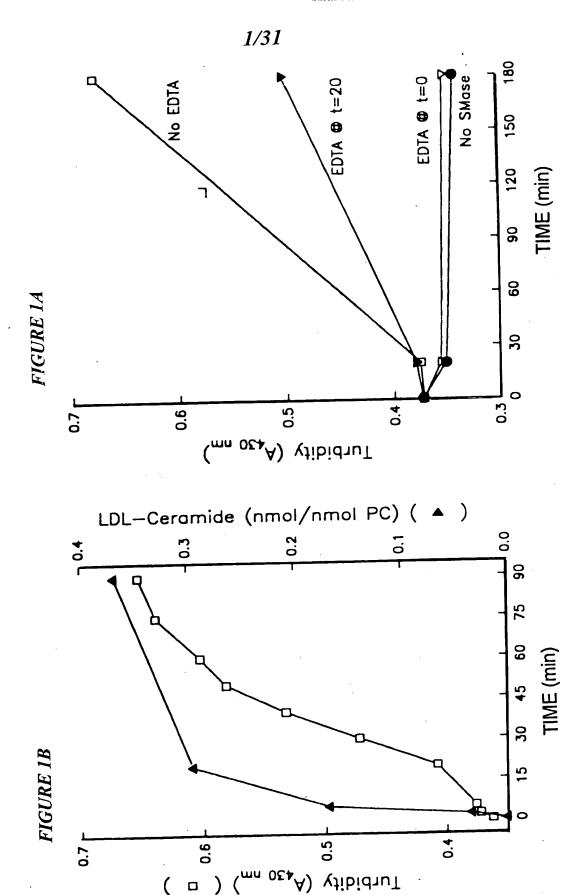
Applicants: Ira Tabas, et al. U.S. Serial No.: 09/518,805 Filed: March 3, 2000

Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc Sphingomyelinase

Sheets 1 of 31



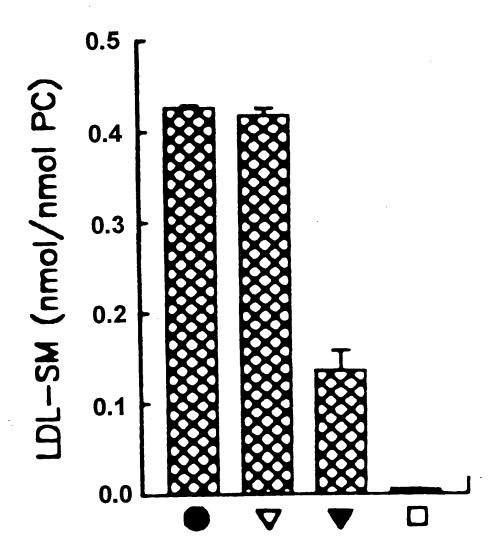
Applicants: Ira Tabas, et al. U.S. Serial No.: 09/518,805

Filed: March 3, 2000 Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc Sphingomyelinase

Sheets 2 of 31

2/31

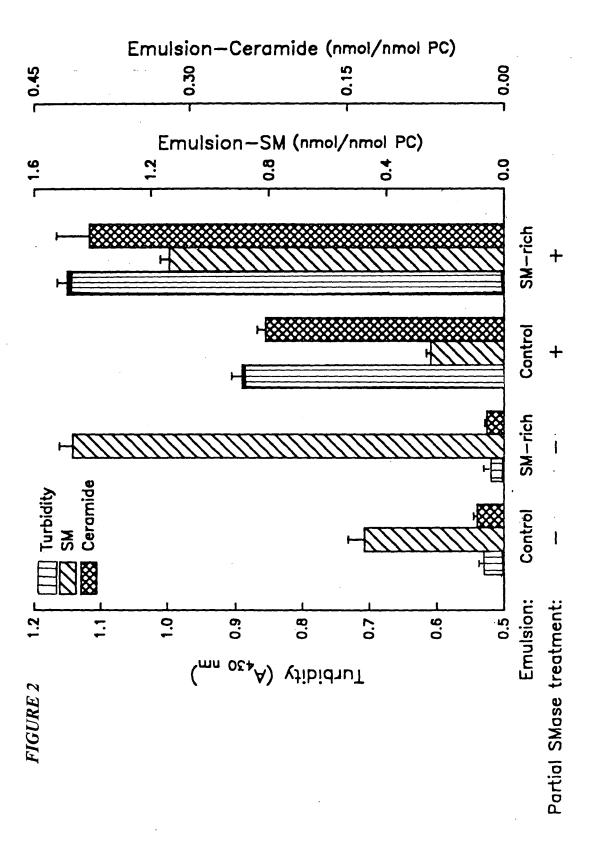
FIGURE 1C



Applicants: Ira Tabas, et al.
U.S. Serial No.: 09/518,805
Filed: March 3, 2000
Title: Method For Treating A Subject Suffering From
Conditions Associated with An Extracellular Zinc
Sphingomyelinase

3/31

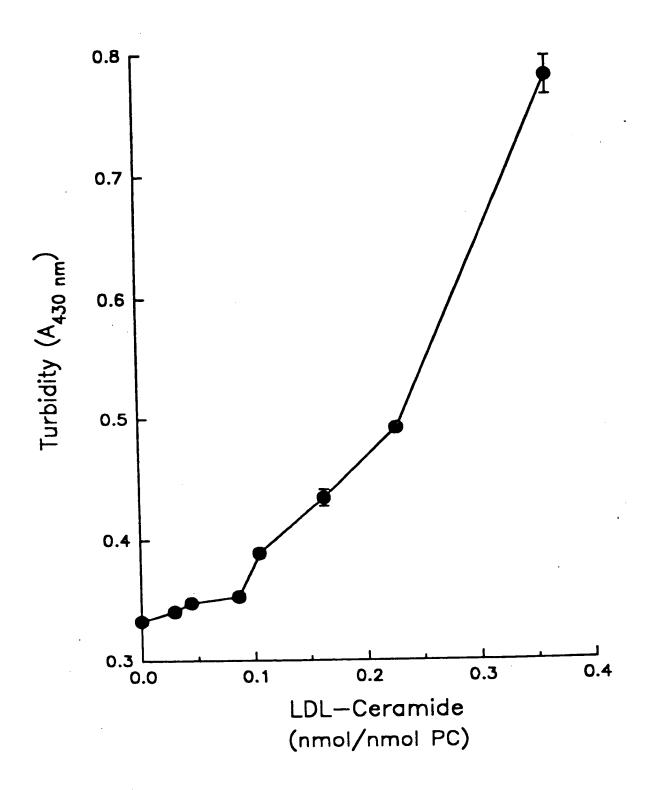
Sheets 3 of 31



Applicants: Ira 1 abas, et al.
U.S. Serial No.: 09/518,805
Filed: March 3, 2000
Title: Method For Treating A Subject Suffering From
Conditions Associated with An Extracellular Zinc
Sphingomyelinase
Sheets 4 of 31

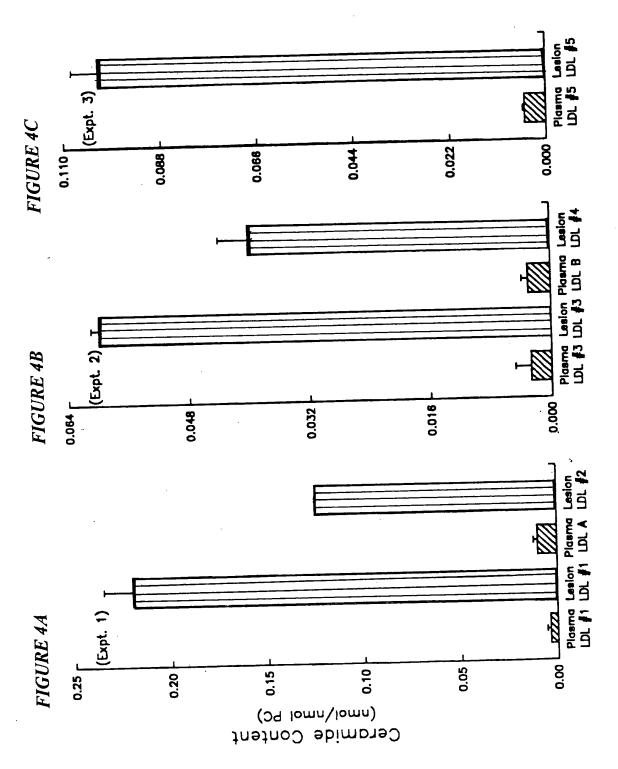
Sheets 4 of 31

FIGURE 3



Sheets 5 of 31

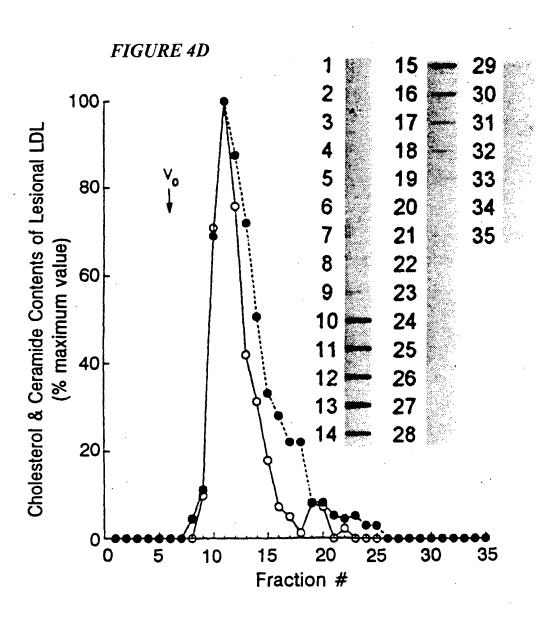




Applicants: Ira Tabas, et al. U.S. Serial No.: 09/518,805 Filed: March 3, 2000

Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc Sphingomyelinase

Sheets 6 of 31

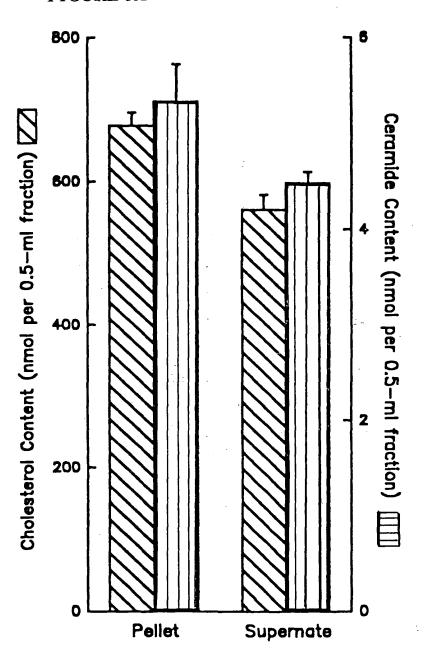


Applicants: Ira Tabas, et al.
U.S. Serial No.: 09/518,805
Filed: March 3, 2000
Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc Sphingomyelinase
Sheets 7 of 31

Sheets 7 of 31

7/31

FIGURE 5A



Applicants: Ira Tabas, et al.
U.S. Serial No.: 09/518,805
Filed: March 3, 2000
Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc Sphingomyelinase

8/31

Sheets 8 of 31



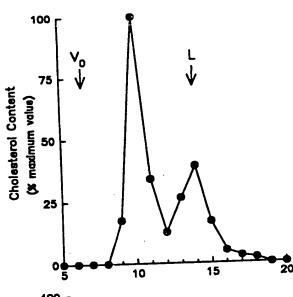


FIGURE 5C

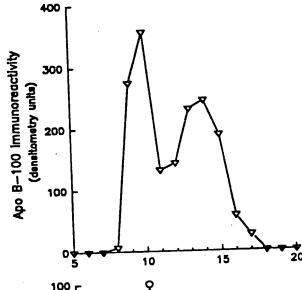
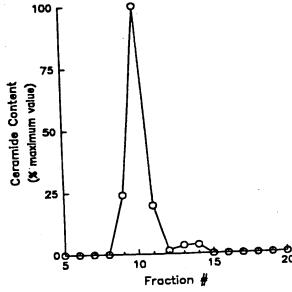
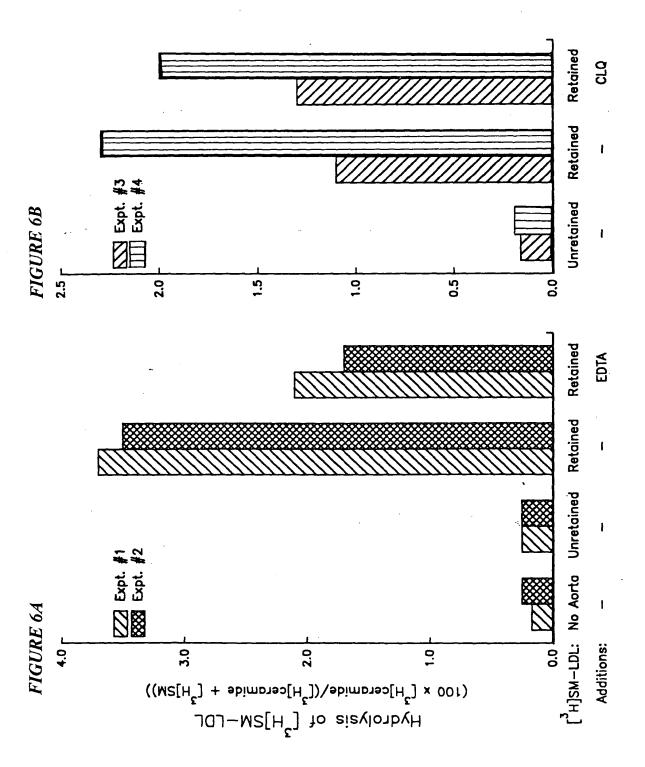


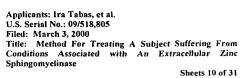
FIGURE 5D

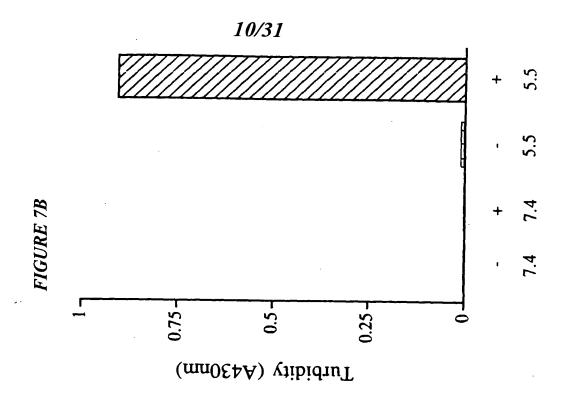


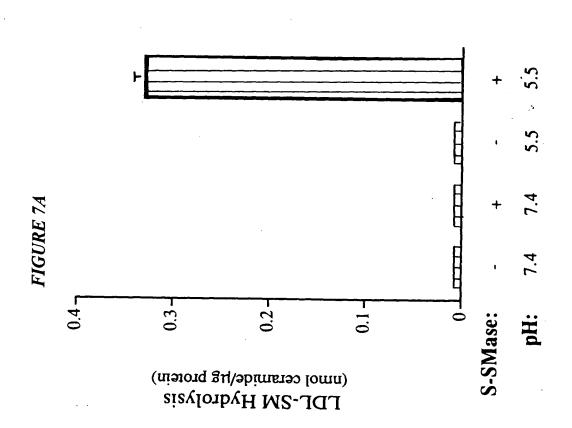
Applicants: 1ra Tabas, et al. U.S. Serial No.: 09/518,805 U.S. Serial No.: 09/518,805
Filed: March 3, 2000
Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc Sphingomyelinase

Sheets 9 of 31





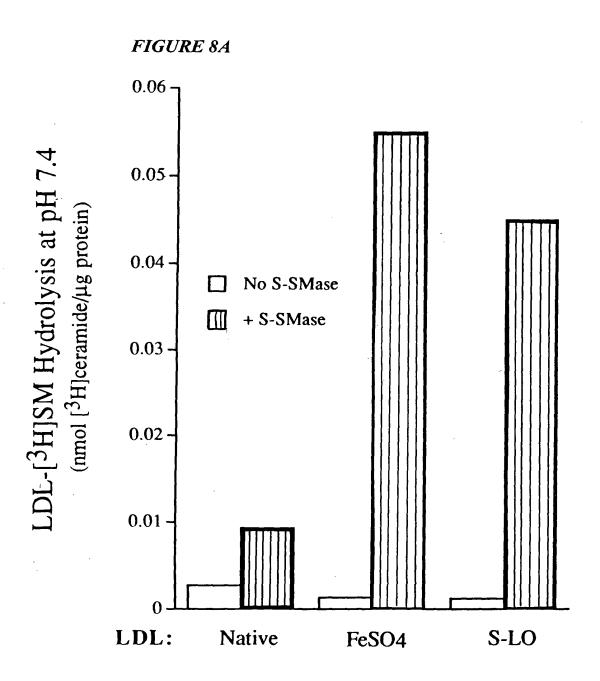




Applicants: Ira Tabas, et al. U.S. Serial No.: 09/518,805

Ed., Section 1802, 097518,005 Filed: March 3, 2000 Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc Sphingomyelinase

Sheets 11 of 31



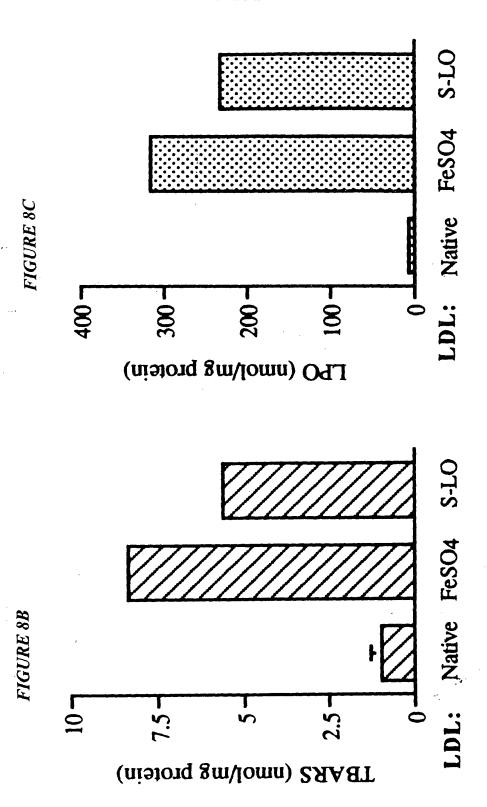
Applicants: Ira Tabas, et al. U.S. Serial No.: 09/518,805 Filed: March 3, 2000

Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc

Sphingomyelinase

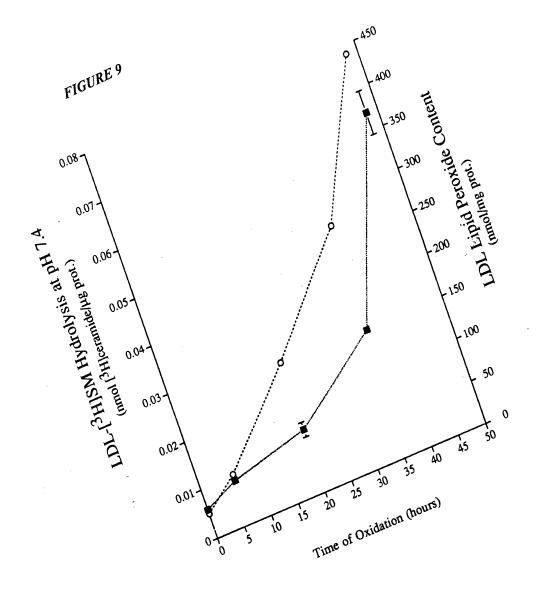
Sheets 12 of 31







13/31

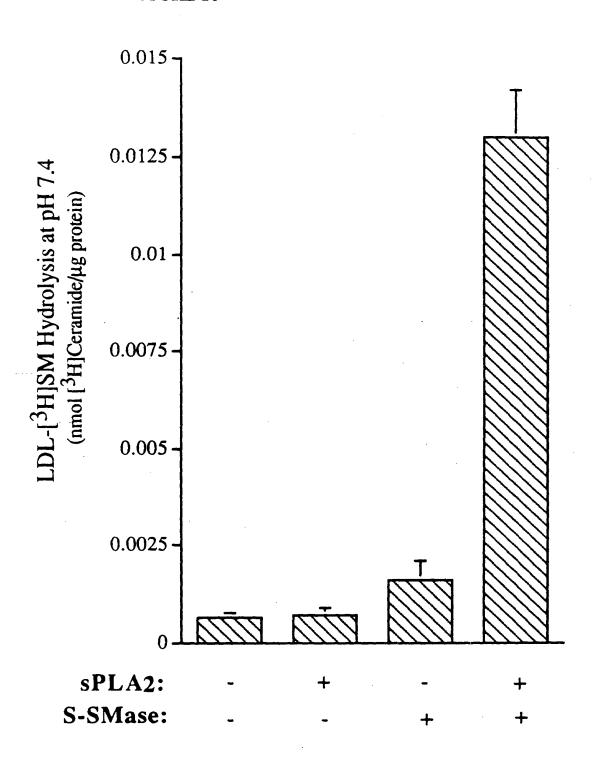


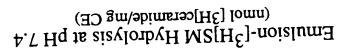
Applicants: Ira Tabas, et al.
U.S. Serial No.: 09/518,805
Filed: March 3, 2000
Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc Sphingomyelinase

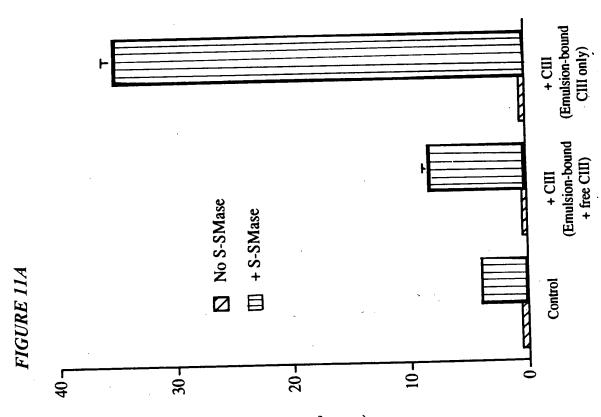
Sheets 14 of 31

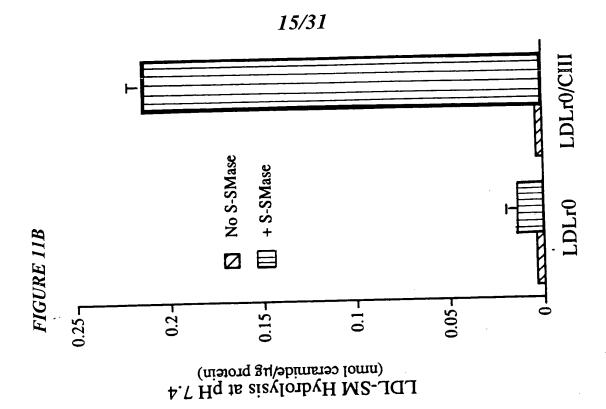
14/31

FIGURE 10









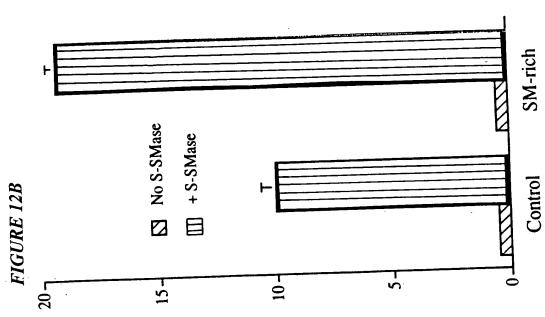
Sphingomyelinase Sheets 15 of 31

Applicants: Ira Tabas, et al. U.S. Serial No.: 09/518,805 Filed: March 3, 2000 Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc Applicants: Ira Tabas, et al. U.S. Serial No.: 09/518,805

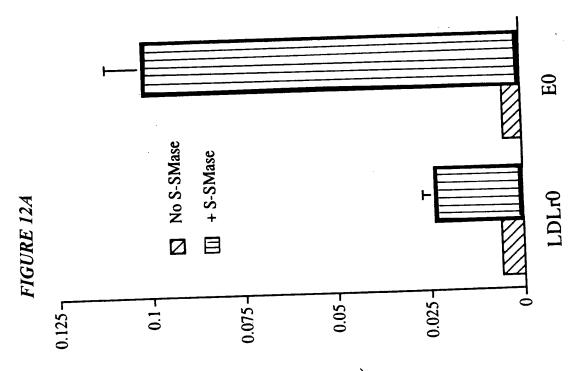
Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc Sphingomyelinase Filed: March 3, 2000

Sheets 16 of 31





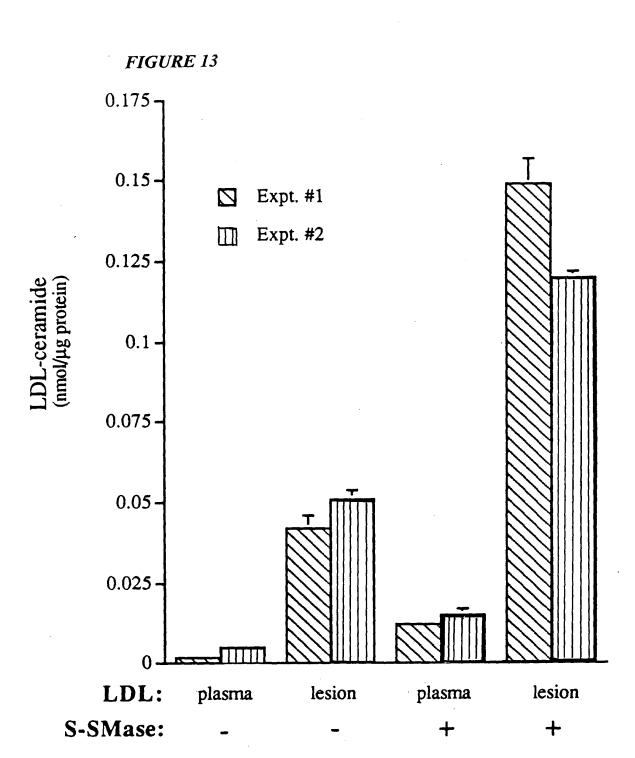
Emulsion- $[^3H]SM$ Hydrolysis at pH 7.4 (nmol $[^3H]$ ceramide/mg CE)

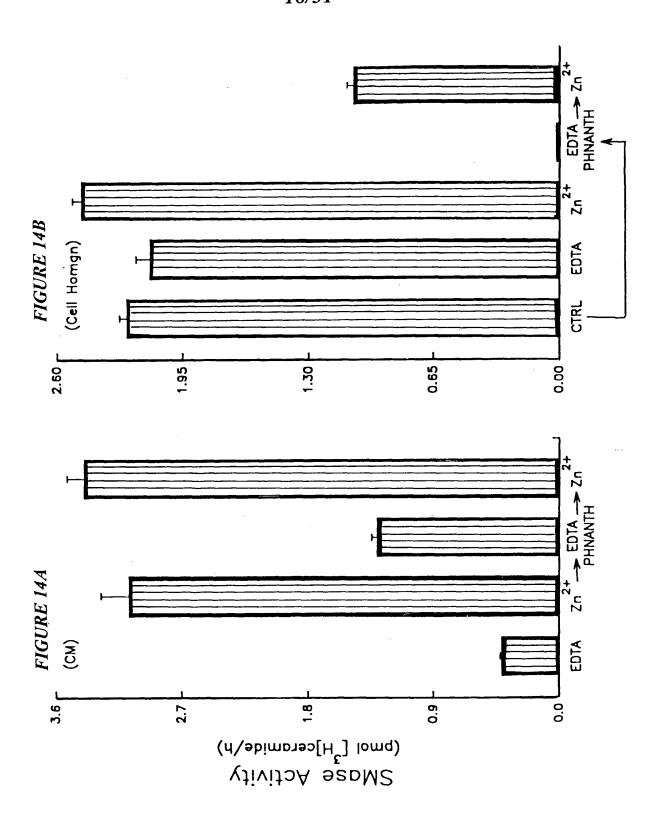


(nmol ceramide/µg protein) LDL-SM Hydrolysis at pH 7.4

Applicants: Ira Tabas, et al.
U.S. Serial No.: 09/518,805
Filed: March 3, 2000
Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc Sphingomyelinase

Sheets 17 of 31





18/31

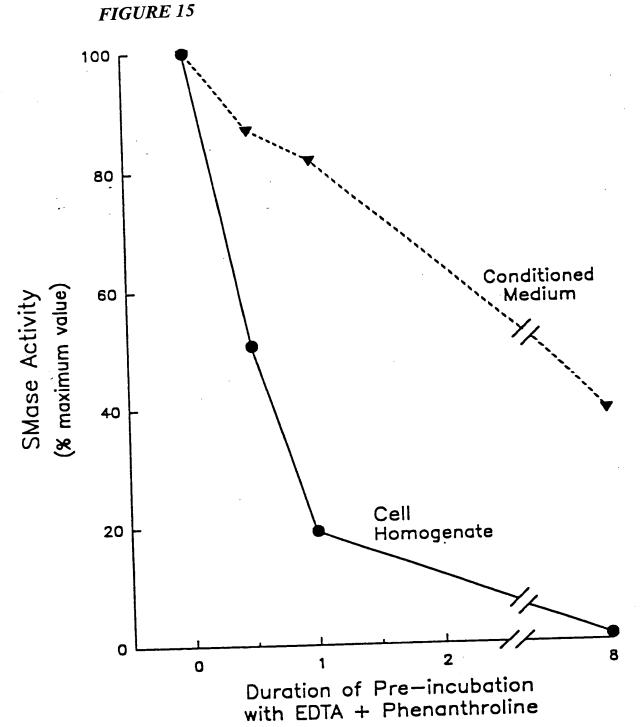
Applicants: Ira Fabas, et al.
U.S. Serial No.: 09/518,805
Filed: March 3, 2000
Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc Sphingomyelinase
Sheets 18 of 31

Applicants: Ira Tabas, et al. U.S. Serial No.: 09/518,805 Filed: March 3, 2000

Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc Sphingomyelinase

Sheets 19 of 31



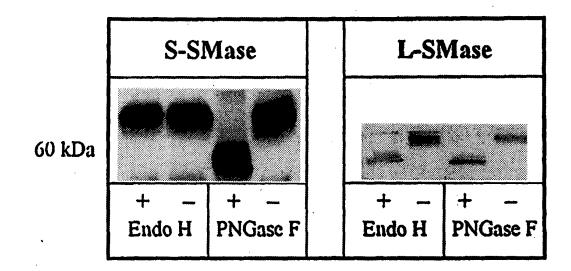


Applicants: Ira Tabas, et al.
U.S. Serial No.: 09/518,805
Filed: March 3, 2000
Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc Sphingomyelinase

Sheets 20 of 31

20/31

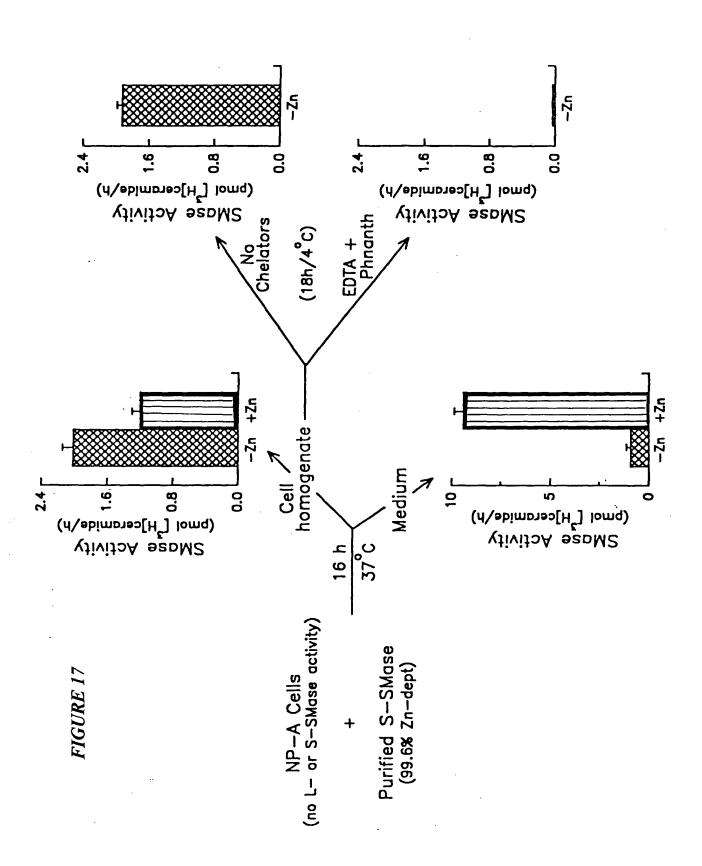
FIGURE 16

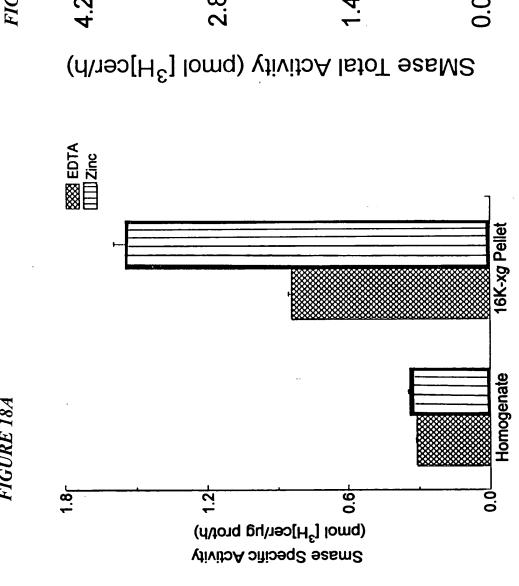


Applicants: fra labas, et al.
U.S. Serial No.: 09/518,805
Filed: March 3, 2000
Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc

Sphingomyelinase

Sheets 21 of 31



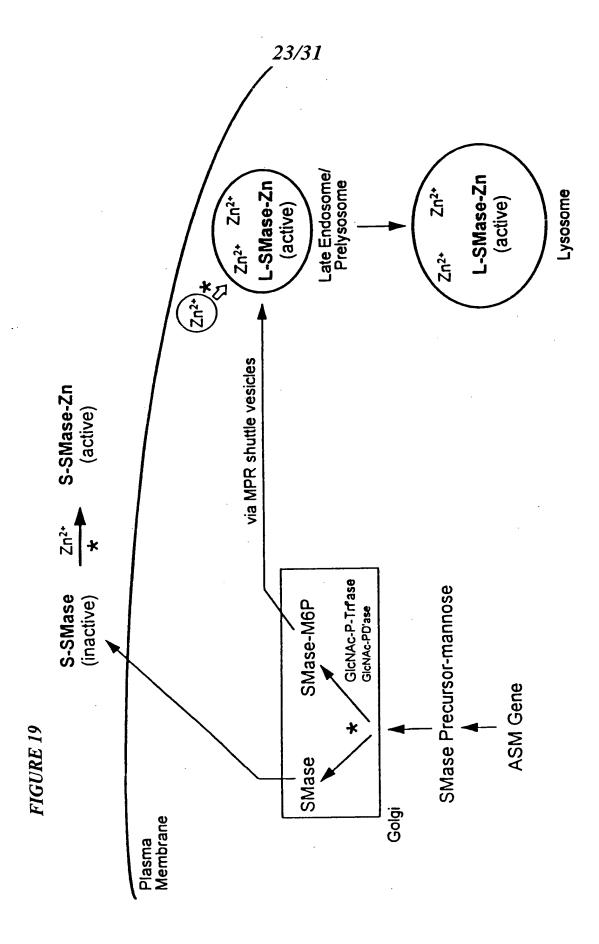


Applicants: 1ra Tabas, et al.
U.S. Serial No.: 09/518,805
Filed: March 3, 2000
Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc Sphingomyelinase

Sheets 22 of 31

Applicants: Ira Tabas, et al.
U.S. Serial No.: 09/518,805
Filed: March 3, 2000
Title: Method For Treating A Subject Suffering From
Conditions Associated with An Extracelletar Zinc Sphingomyelinase

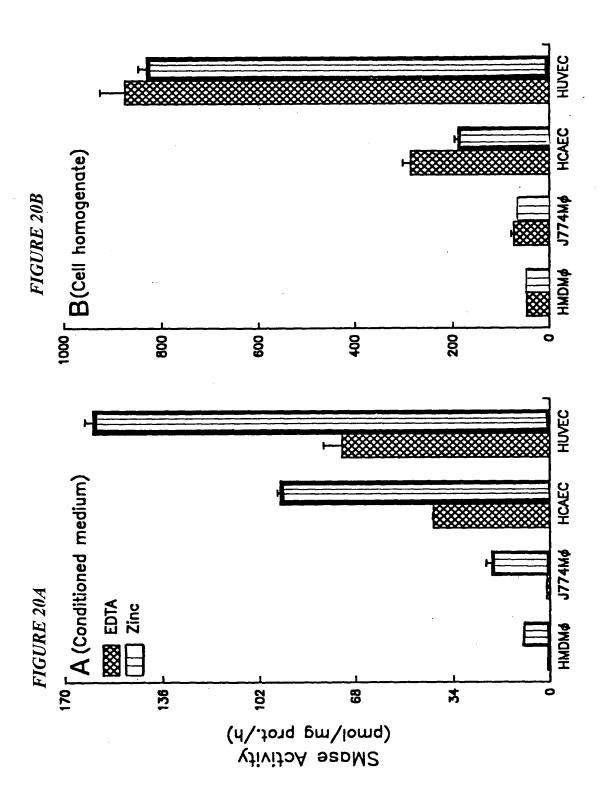
Sheets 23 of 31



Applicants. Ira Tabas, et al
U.S. Serial No.: 09/518,805
Filed: March 3, 2000
Title: Method For Treating A Subject Suffering From
Conditions Associated with An Extracellular Zinc
Sphingomyelinase
Sheets 24 of 31

Sheets 24 of 31

24.′31



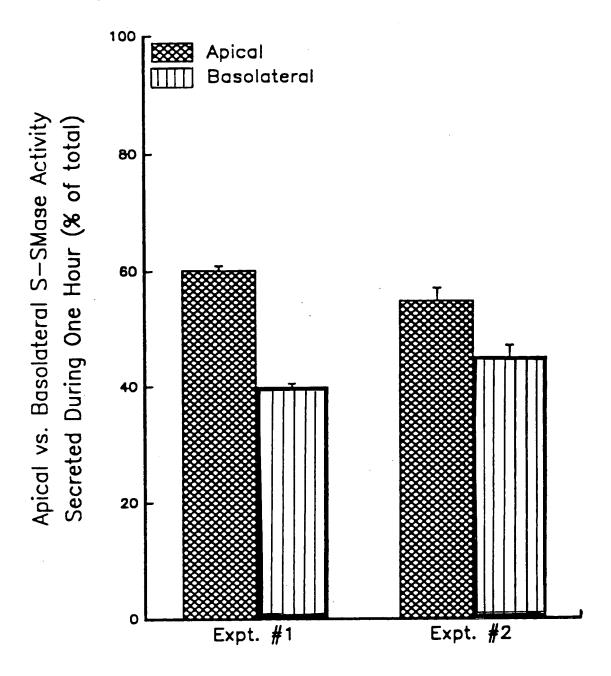
Applicants: Ira Tabas, et al. U.S. Serial No.: 09/518,805 Filed: March 3, 2000 Title: Method For Treating A Subject

Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc Sphingomyelinase

Sheets 25 of 31

25/31

FIGURE 21A



Applicants: Ira Tabas, et al. U.S. Serial No.: 09/518,805 Filed: March 3, 2000

Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc

Sphingomyelinase

Sheet 26 of 31

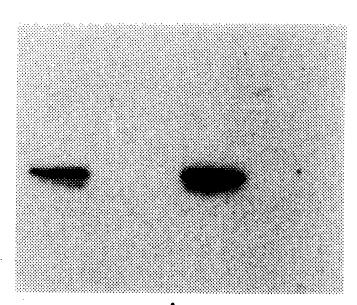
t orania establica transcribente en establica establica en establica en establica en el constitución de la con

26/31

FIGURE 21B

148 kD

60 kD



ap bl ap bl Expt. #2

U.S. Serial (NO.: 09/510,005)
Filed: March 3, 2000
Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc Sphingomyclinase

Sheets 27 of 31

27/31

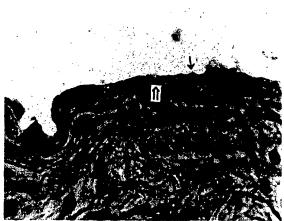
FIGURE 22A

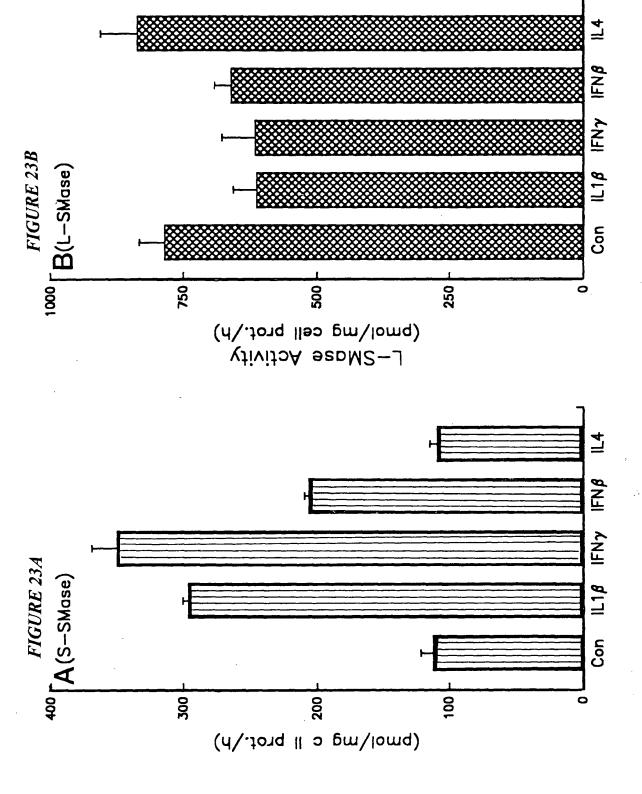
FIGURE 22B

A (SMase Knockout)



B (Wild type)





28/31

Applicants: Ira Tabas, et al. U.S. Serial No.: 09/518,805 C.S. Serial (w. 1973) (1973)
Filed: March 3, 2000
Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc Sphingomyelinase

Shorts 28 of 31

Sheets 28 of 31

C.S. Serial Nu.: 09/518,805 Filed: March 3, 2000 Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc Sphingomyclinase Sheets 2 of 31

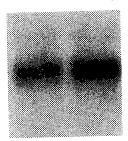
Sheets 2 of 31

29/31

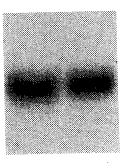
FIGURE 24

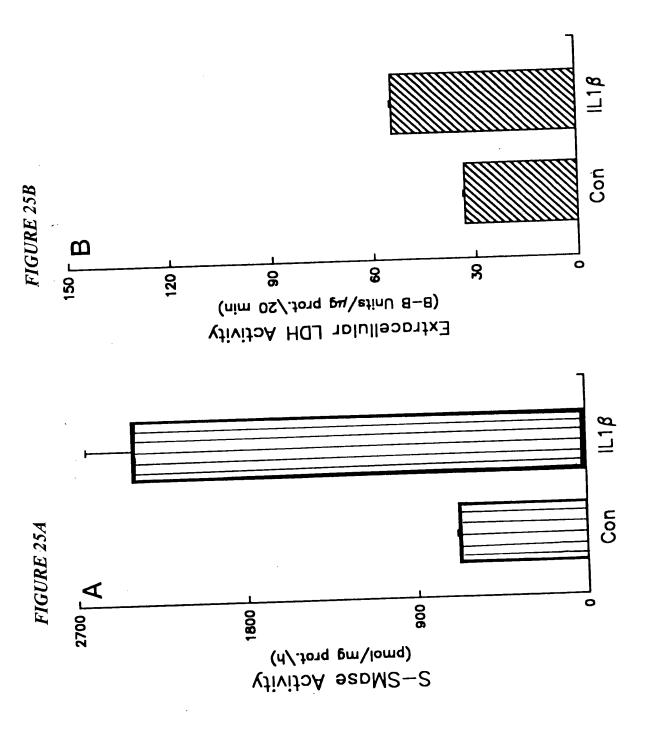
Con IL1B





 $GAPDH \rightarrow$





30/31

Applicants, Ira Tabas, et al
U.S. Serial No.: 09/518,805
Filed: March 3, 2000
Title: Method For Treating A Subject Suffering From
Conditions Associated with An Extracellular Zinc
Sphingomyelinase

Applicants: Ira Tabas, et al. U.S. Serial No.: 09/518,805

Filed: March 3, 2000

Title: Method For Treating A Subject Suffering From Conditions Associated with An Extracellular Zinc Sphingomyelinase

31/31

FIGURE 26

